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Creation of a Novel Algorithm to Identify Patients with Becker and Duchenne Muscular Dystrophy within an Administrative Database and Application of the Algorithm to Assess Cardiovascular Morbidity

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Abstract

Background—Outcome analyses in large administrative databases are ideal for rare diseases such as Becker and Duchenne muscular dystrophy. Unfortunately, Becker and Duchenne do not yet have specific ICD-9/ICD-10 codes. We hypothesized that an algorithm could accurately identify these patients within administrative data and improve assessment of cardiovascular morbidity.

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CONFLICTS OF INTEREST

None

ETHICAL STANDARDS

The authors assert that all procedures contributing to this work comply with the ethical standards of the relevant national guidelines on human experimentation Belmont Report and with the Helsinki Declaration of 1975, as revised in 2008, and has been approved by the institutional committees the Institutional Review Boards of Vanderbilt University Medical Center, Nationwide Children's Hospital, and Children's National Medical Center approved this study.

Methods—Hospital discharges (n=13,189) for patients with muscular dystrophy (ICD-9 359.1) were identified in the Pediatric Health Information System database. An identification algorithm was created then validated at 3 institutions. Multivariable generalized linear mixed effects models used to estimate the associations of length of stay, hospitalization cost, and 14-day readmission with age, encounter severity, and respiratory disease accounting for clustering within a hospital.

Results—The identification algorithm improved identification of Becker and Duchenne patients from 55% (359.1 alone) to 77%. On bivariate analysis, left ventricular dysfunction and arrhythmia were associated with increased cost of hospitalization, length of stay, and mortality ($p<0.001$). After adjustment, Becker and Duchenne patients with left ventricular dysfunction and arrhythmia had increased length of stay (rate ratio 1.4 and 1.2, $p<0.001$ and $p=0.004$) and cost (rate ratio 1.4 and 1.4, both $p<0.001$).

Conclusions—Our algorithm accurately identifies patients with Becker and Duchenne and can be used for future analysis of administrative data. Our analysis demonstrates the significant effects of cardiovascular disease on length of stay and hospitalization cost in Becker and Duchenne patients. Better recognition of the contribution of cardiovascular disease during hospitalization with earlier, more intensive evaluation and therapy may help improve outcomes in this patient population.

Keywords

Duchenne muscular dystrophy; Becker muscular dystrophy; Pediatric Health Information System; Left ventricular dysfunction; Arrhythmia; Cardiomyopathy

1. INTRODUCTION

Becker and Duchenne muscular dystrophy are the most common muscular dystrophies, with an incidence of 1 in 18500 and 1 in 4700 live male births, respectively.^{1, 2} Patients with Becker and Duchenne muscular dystrophy have a mutation in the DMD gene that leads to a deficiency or absence of dystrophin protein.^{3, 4} This results in skeletal muscle weakness and loss of ambulation, usually in the second decade of life in Duchenne muscular dystrophy and later in life in the milder Becker muscular dystrophy. Both reduced and absent dystrophin also leads to the development of cardiomyopathy resulting in progressive left ventricular dysfunction.^{5, 6}

Understanding of the clinical importance of Becker and Duchenne cardiomyopathy has increased significantly over the past few decades. This is primarily due to improvements in care, which have increased life expectancy in patients with Duchenne muscular dystrophy and unmasked the cardiovascular phenotype.⁷ The preponderance of Becker and Duchenne research focuses on skeletal muscle weakness, but cardiovascular disease is now one of the leading causes of death in patients with Duchenne muscular dystrophy.⁸ Highlighting the effects of cardiovascular disease on inpatient morbidity (length of stay, cost of hospitalization, readmission rate) in patients with Becker and Duchenne may help improve recognition of the problems associated with cardiovascular disease. Moreover, identifying risk factors leading to increased length of stay, cost of hospitalization, and readmission rates

in this population can increase recognition and potentially improve outcomes in future admissions.

Because Becker and Duchenne muscular dystrophy are relatively rare, the true impact of cardiovascular disease can be difficult to determine. Research using large administrative databases can overcome these issues. However, using the International Classification of Disease (ICD)-9 billing codes, Becker and Duchenne muscular dystrophy are coded as 359.1, a non-specific code that also encompasses multiple other neuromuscular diagnoses (Table 1). This non-specific coding can make analysis problematic. Indeed, a recent manuscript using the Pediatric Health Information System database to evaluate the effect of cardiovascular disease in Duchenne muscular dystrophy was limited by this non-specific coding.⁹ We hypothesized that, by creating a novel list of exclusion criteria which could be validated, patients with Becker and Duchenne muscular dystrophy could be more accurately identified within an administrative and billing database, allowing for the current study and future assessments in this patient population. To demonstrate the potential utility of this technique, we used the Pediatric Health Information System database to assess cardiovascular risk factors associated with increased length of stay, cost of hospitalization, and 14-day readmission in patients with Becker and Duchenne muscular dystrophy.

2. MATERIALS AND METHODS

2.1 Pediatric Health Information System Database and Identification Algorithm

This multicenter retrospective cohort study utilized administrative data from the Pediatric Health Information System database. The database contains inpatient, emergency department, ambulatory surgery, and observation encounters from 49 tertiary children's hospitals in the United States. Data are de-identified at the time of submission and are subjected to reliability and validity checks by participating hospitals, Children's Hospital Association (Lenexa, KS), and Truven Health Analytics (Ann Arbor, MI) before inclusion in the database. Patients are assigned up to 41 ICD-9/10 diagnosis and procedure codes, and consistently encrypted medical record numbers allow longitudinal tracking of patients across encounters to the same hospital. The Pediatric Health Information System database represents approximately 15% of the national pediatric hospitalizations and 46.4% of children's hospitals' total volume.

The Institutional Review Boards of Vanderbilt University Medical Center, Nationwide Children's Hospital, and Children's National Medical Center approved this study. Using ICD-9 diagnosis codes (conversion to ICD-10 codes occurred in the last quarter of 2015), an algorithm of exclusion criteria termed the "identification algorithm" was created to improve the identification of patients with Becker and Duchenne muscular dystrophy. The effectiveness of the algorithm was evaluated by "re-identification" of patients from three sites. The electronic medical record was queried to determine the diagnosis and gender of all patients with an ICD-9 code of 359.1 and the diagnosis of the remaining patients after the identification algorithm was applied.

The algorithm was constructed to optimize the specificity of identifying patients with Becker and Duchenne muscular dystrophy. Given the size of the Pediatric Health Information

System database, investigators determined that incorrectly excluding patients with Becker and Duchenne muscular dystrophy was preferable to incorrectly including patients with an alternate diagnosis. The goal at the outset of the study was to construct an algorithm where at least 80% of remaining patients carried a diagnosis of Becker and Duchenne muscular dystrophy. All patients with an ICD-9 of 359.1 were initially included. In order to eliminate errors in coding, patients with a change in primary diagnosis to a different 359.x code were excluded. A set of clinical characteristics was created to exclude neuromuscular conditions other than Becker and Duchenne muscular dystrophy. Briefly, patients with severe disease at a young age felt unlikely to be due to Becker or Duchenne were eliminated (early mortality, ventilatory support, or cardiovascular disease). Because the majority of Becker and Duchenne patients do not require gastrostomy tubes until later in life, and only 15% of patients were older than 18 in this cohort, all patients with gastrostomy tubes were removed. Other neurologic diagnoses that could have been mistakenly coded as 359.1 were removed, as were diagnoses that came up in an initial search of 359.1 at one institution. Patients undergoing heart transplantation were not excluded as transplantation has been performed in the Becker muscular dystrophy population.

2.2 Primary Analysis

The primary analysis was performed on the entire cohort (49 hospitals) after application of the identification algorithm. Discharges in patients with muscular dystrophy were identified in Pediatric Health Information System from January 1, 2003 - September 30, 2015 (chosen based on transition from ICD-9 to ICD-10). Patients with left ventricular dysfunction and arrhythmia were identified using ICD-9 codes (Supplementary Table S1). Patients requiring respiratory support (Table S1) were also identified. Discharges that occurred under 1 year of age were eliminated from the analysis as these admissions were more likely to reflect neonatal and infant hospitalizations unrelated to the diagnosis of Becker and Duchenne muscular dystrophy. The primary outcome measures included length of stay, cost of hospitalization (estimated from charges using hospital/year specific ratios of cost to charge), 14-day all-cause readmission, and in-hospital mortality.

2.3 Statistical Analysis

To assess the effectiveness of the identification algorithm, the percentage of the patients with Becker and Duchenne muscular dystrophy before and after application of the algorithm was calculated for the 3 sites. The percentage of patients with Becker and Duchenne muscular dystrophy who were incorrectly excluded was also calculated.

For the primary analysis on the entire Pediatric Health Information System cohort, pertinent demographic information was summarized and bivariate analyses were performed to assess the relationships of each demographic and length of stay (stratified by <2 days vs. 2+ days) using chi-square and Wilcoxon Rank Sum tests. Next, separate generalized linear mixed effects models assuming either an exponential distribution (length of stay and cost) or a binomial distribution (14-day readmission and mortality) for the outcome were used to estimate the adjusted associations with age, encounter severity, and respiratory disease. A random intercept was used to account for correlation arising from taking repeated measurements on the same hospital. Results are summarized using the rate ratio or odds

ratio with corresponding 95% confidence intervals. Statistical analyses were performed using SAS v.9.4 (SAS Institute, Cary, NC), and a p-value < 0.05 was considered statistically significant.

3. RESULTS

3.1 Identification Algorithm

The final identification algorithm is listed in Table 2. At the 3 Pediatric Health Information System sites, we performed a detailed assessment of our algorithm by re-identifying the 289 patients with an ICD-9 of 359.1 and determining true diagnosis using the local electronic medical record. Of the 289 patients, 55% (158) had a diagnosis of Becker and Duchenne muscular dystrophy prior to the application of the identification algorithm. Excluding female patients increased this to 68% (158 of 233). Applying the identification algorithm increased the accuracy of correct Becker and Duchenne muscular dystrophy diagnosis to 77% (131 of 170) by excluding 27 patients with Becker and Duchenne muscular dystrophy (17%) and 91 patients without Becker and Duchenne muscular dystrophy (70% of non-Becker and non-Duchenne muscular dystrophy patients). Of note, two sites provided a breakdown of Becker muscular dystrophy vs Duchenne muscular dystrophy, and only 5% of included patients at those sites had Becker muscular dystrophy. We hypothesize that this preponderance of Duchenne muscular dystrophy is due to the relatively higher frequency in the general population and earlier cardiomyopathy as well as a lower frequency of admission in children with Becker muscular dystrophy.

3.2 Demographics

In the entire Pediatric Health Information System cohort, a total of 3,430 unique patients with 13,189 discharges from 49 hospitals were identified with an ICD-9 of 359.1. After application of the identification algorithm, 1,916 patients and 4014 discharges remained.

The majority of discharges (89.5%) were to home, 4% were discharged to a skilled facility, 3.1% to home health service, and 2.2% (90 discharges) died in the hospital (4.7% of patients). As expected, neuromuscular and cardiovascular codes were the most common, with 90% of discharges having at least one neuromuscular code and 30% having at least one cardiovascular code. In addition, 22% had a code for congenital or genetic defects, 21% had a code suggesting technology dependence, and 7% had a code denoting respiratory disease. A total of 26% of discharges had a diagnosis of left ventricular dysfunction, 8% arrhythmia, and 21% respiratory disease. Further demographic data are reported in Table 3.

3.3 Primary Analysis

Bivariate analysis for the predictors left ventricular dysfunction, arrhythmia, and respiratory disease for the outcomes of length of stay, cost of hospitalization, 14-day readmission, and mortality are shown in Table 4. We also considered adjusted models controlling for age, encounter severity, respiratory disease, and hospital. The adjusted association of predictors with mortality was not estimated because there were too few events to support a multivariable model. However, for length of stay we found significant adjusted association with left ventricular dysfunction (rate ratio = 1.4, 95% CI [1.3, 1.5], p<0.001), arrhythmia

(rate ratio = 1.2, 95% CI [1.1, 1.4], $p=0.004$), and respiratory disease (rate ratio = 1.6, 95% CI [1.5, 1.8], $p<0.001$) (Table 5). We also found significant adjusted association for cost of hospitalization with left ventricular dysfunction (rate ratio = 1.2, 95% CI [1.1, 1.3], $p<0.001$), arrhythmia (rate ratio = 1.4, 95% CI [1.2, 1.6], $p<0.001$), and respiratory disease (rate ratio = 1.4, 95% CI [1.3, 1.5], $p<0.001$). Becker and Duchenne muscular dystrophy patients with respiratory disease had decreased rates of 14-day readmission (odds ratio = 0.6, 95% CI [0.3, 1], $p=0.045$), while left ventricular dysfunction (odds ratio = 1, 95% CI [0.7, 1.3], $p=0.780$) and arrhythmia (odds ratio = 1.5, 95% CI [0.9, 2.5], $p=0.128$) were not significantly associated with readmission. The combination of arrhythmia and left ventricular dysfunction (cardiovascular disease) demonstrated similar effects for hospitalization cost and length of stay (Supplementary Table S2).

4. DISCUSSION

These results demonstrate that: 1) the application of our identification algorithm confirmed by direct validation significantly increases the specificity for Becker and Duchenne muscular dystrophy in the Pediatric Health Information System database and 2) left ventricular dysfunction, arrhythmia, and respiratory disease play significant, independent roles in predicting inpatient morbidity and, in the bivariate model, mortality. Given the rarity of Becker and Duchenne muscular dystrophy, a method that allows researchers to leverage large databases is critical to improving outcomes. Analysis of the Pediatric Health Information System database may help identify areas of improvement that could lead to a decrease in the costs of hospitalization for patients with Becker and Duchenne muscular dystrophy. Moreover, these data can serve as a baseline appraisal, allowing for the assessment of future shifts in outcome resulting from modifications in therapies or changes to standards of care in Becker and Duchenne muscular dystrophy. To allow for future analyses, we have translated our exclusion criteria to ICD-10 diagnostic codes (Supplementary Table S3).

The ICD-9 billing code that includes Becker and Duchenne muscular dystrophy also includes many other diagnoses. Fortunately, this will be rectified in October of 2018 with a new ICD-10 code. However, given that this manuscript used over 12 years of data from the Pediatric Health Information System, it will take an extended period of time for that change to be relevant. The algorithm reported here will remain critical for future analyses and will facilitate analysis of historical trends. In addition, similar methods can be used to evaluate morbidity and mortality due to other forms of muscular dystrophy coded as 359.1.

These data further emphasize the importance of cardiovascular disease in Becker and Duchenne muscular dystrophy. It is notable that 30% of the patients in this analysis had at least one cardiovascular code while only 7% had a code for respiratory disease. Cardiovascular disease has become increasingly recognized as a major contributor of mortality.^{8,10} Punnoose et al recently evaluated the association between cardiovascular disease and morbidity and mortality in the Pediatric Health Information System database, though their analysis was limited by the poor specificity of ICD-9 coding for Duchenne muscular dystrophy.⁹ Their study also demonstrated that ventricular tachycardia and heart failure, as well as chronic ventilator use, were risk factors for the combined outcome of

cardiac arrest or death. Their exclusions likely improved specificity for Duchenne muscular dystrophy but are less extensive than the algorithm we present here, and they were unable to directly validate the effectiveness of their exclusion criteria. The prevalence of cardiomyopathy increases significantly as Becker and Duchenne muscular dystrophy patients age, with previous studies suggesting that >60% of patients over 18 years of age have cardiomyopathy.^{5, 11} Unfortunately, while official recommendations state that echocardiographic screening should begin by 6 years of age, Spurney et al demonstrated that a surprisingly large portion of Duchenne muscular dystrophy patients had not undergone echocardiography by 10 years of age.¹¹ Moreover, that study revealed that over half of Duchenne muscular dystrophy patients with a diagnosis of cardiomyopathy were not prescribed appropriate anti-congestive therapy. Taken together, these data suggest that the focus of care in Becker and Duchenne muscular dystrophy must shift towards earlier detection and prevention of cardiovascular disease. Raising awareness of the risks of cardiovascular disease in Becker and Duchenne muscular dystrophy patients admitted to the hospital may lead to earlier recognition and more aggressive therapy, eventually improving outcomes.

Our data confirm that respiratory disease continues to play a significant role in inpatient morbidity and mortality in patients with Becker and Duchenne muscular dystrophy. Multiple authors have reported the improved life expectancy associated with ventilation in Duchenne muscular dystrophy.^{7, 8, 12} While survival has increased significantly with these interventions, respiratory disease of any kind, including acute respiratory complications and the need for chronic ventilation, significantly increased the odds of a longer length of stay and higher cost of hospitalization. Moreover, the complex interactions between cardiovascular and respiratory disease likely plays an important role in Becker and Duchenne muscular dystrophy inpatient morbidity.

Multiple reports have demonstrated the increased cost of both Duchenne and Becker muscular dystrophy compared with the general population. Thayer et al demonstrated a 10-fold increase in costs in a small cohort of 75 Duchenne muscular dystrophy patients by analyzing claims from a single health plan.¹³ Surveys of families in multiple countries have demonstrated increases in both healthcare costs and out of pocket costs to families of Becker and Duchenne muscular dystrophy patients.^{14–16} In general, costs increase with increasing age/progression of disease.^{14, 15} Costs have also been shown to be increased in Duchenne muscular dystrophy patients with tracheostomy compared to those on continuous non-invasive ventilation.¹⁷ However, only our data and that of Punnoose have stratified costs based on cardiovascular disease.⁹ Awareness of the increased cost associated with both arrhythmia and left ventricular dysfunction could allow for improvement in Becker and Duchenne muscular dystrophy costs with earlier and more aggressive detection and therapy. Not surprisingly, our data also demonstrated significantly increased length of stay and cost of hospitalization with advancing age in Becker and Duchenne muscular dystrophy, even when correcting for arrhythmia, left ventricular dysfunction, and respiratory disease.

4.1 Limitations

This study does have inherent limitations. As with any large database, data could be incomplete, incorrect, or missing. The Pediatric Health Information System does not capture events that occur outside a hospital encounter and does not include physician fees as part of the cost analysis. In addition, the Pediatric Health Information System only captures events that occur in children's hospitals. Specific to this study, our algorithm prioritized identification of Becker and Duchenne muscular dystrophy patients with the knowledge that some patients would be incorrectly excluded. However, re-identification demonstrated that 17% of Becker and Duchenne muscular dystrophy patients were incorrectly excluded while 70% of non-Becker and Duchenne muscular dystrophy were excluded. The algorithm we present to better identify Becker and Duchenne muscular dystrophy patients does still include patients without Becker and Duchenne muscular dystrophy, but this number is significantly lower after our exclusion criteria and we feel the advantages of using larger numbers for this and future studies will outweigh this limitation. We also evaluated an algorithm that increased the identification of Becker and Duchenne muscular dystrophy patients to 79% (Supplementary Table 4), but this algorithm excluded a much larger number of patients with Becker's and Duchenne's (35%), so the less aggressive algorithm was used for the final analysis. Of note, the results of the primary analysis were not significantly different between algorithms.

4.2 Conclusions

We present an algorithm for improving identification of Becker and Duchenne muscular dystrophy patients using an administrative database. Initial analysis demonstrated a large number of Becker and Duchenne muscular dystrophy patients with cardiovascular codes and increased cost and length of stay in patients with both arrhythmia and left ventricular dysfunction. This algorithm can be used to analyze other outcomes of interest in patients with Becker and Duchenne muscular dystrophy and can also serve as a baseline to assess changes in therapy over time.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Table 1:

List of Diagnoses Included in ICD-9 Code 359.1

Facioscapulohumeral muscular dystrophy	Hereditary, progressive muscular dystrophy
Landouzy-Déjérine	Leyden-Möbius
Pseudohypertrophic muscular dystrophy	Distal muscular dystrophy
Duchenne's muscular dystrophy	Facioscapulohumeral muscular dystrophy
Duchenne-Griesinger	Limb-girdle muscular dystrophy
Erb (-Landouzy)	Ocular muscular dystrophy
Hereditary neuromuscular NEC	Oculopharyngeal muscular dystrophy
Duchenne's pseudohypertrophy, muscles	Pelvicurular atrophic muscular dystrophy
Erb's dystrophy	Progressive ophthalmoplegic muscular dystrophy
Gower's muscular dystrophy	Scapuloperoneal ophthalmoplegic muscular dystrophy
Emery Dreifuss muscular dystrophy	Becker's muscular dystrophy
Kearns-Sayre syndrome (disorder)	Mitochondrial ocular myopathy
Muscular dystrophy	Neuropathic muscular dystrophy
Restrictive lung disease due to muscular dystrophy	Congenital myopathy
Congenital muscular dystrophy	

Table 2:**Identification Algorithm**

	Patients Excluded	Patients Remaining
Inclusion		
ICD-9=359.1		3430
Exclusions		
Cerebral degenerations (330.x), Spinocerebellar disease (334.x), Anterior horn cell disease (335.x), or Hereditary and idiopathic peripheral neuropathy (356.x)	387	3043
Changed to another 359.x code	405	2638
G-tube ^a	458	2180
Ophthalmoplegia	2	2178
Pigmentary retinopathy	0	2178
Mortality age<5	8	2170
Ventilation <5 years of age (tracheostomy or CPAP/BIPAP ^b)	0	2170
AICD ^c <10 years of age	1	2169
Arrhythmia <5 years of age	7	2162
Cardiovascular disease <5 years of age	6	2156
Female	224	1932
Age <1 year at time of only encounter	16	1916

^a gastrostomy tube (G-tube)

^b Continuous positive airway pressure (CPAP), bilevel positive airway pressure (BiPAP)

^c Automatic implantable cardioverter-defibrillator (AICD)

Table 3:**Demographics**

		Prior to Exclusions	Post Exclusions
N Discharges		13189	4014
N Patients		3430	1916
Discharge contained ICD-9 code 359.1		7538 (57.2%)	3522 (87.7%)
Age: Median [IQR]		13 [6, 17]	14 [11, 17]
Age	<1	640 (4.9%)	
	1–4	1966 (14.9%)	159 (4%)
	5–9	2105 (16%)	504 (12.6%)
	10–14	3344 (25.4%)	1431 (35.7%)
	15–18	3124 (23.7%)	1293 (32.2%)
	19+	2010 (15.2%)	627 (15.6%)
Gender	Male	9746 (73.9%)	4014 (100%)
Race	Non-Hispanic White	7900 (59.9%)	2590 (64.5%)
	Non-Hispanic Black	1139 (8.6%)	289 (7.2%)
	Hispanic	2826 (21.4%)	730 (18.2%)
	Asian	313 (2.4%)	81 (2%)
	Other	1011 (7.7%)	324 (8.1%)
Discharge Disposition	Home Health Service	692 (5.2%)	123 (3.1%)
	Home	11690 (88.6%)	3593 (89.5%)
	Skilled Facility	385 (2.9%)	159 (4%)
	Died	210 (1.6%)	90 (2.2%)
	Other	212 (1.6%)	49 (1.2%)
CCCs v.2.0	Neuromuscular	11202 (84.9%)	3594 (89.5%)
	Cardiovascular	3189 (24.2%)	1192 (29.7%)
	Disease		
	Respiratory	3248 (24.6%)	272 (6.8%)
	Renal	458 (3.5%)	65 (1.6%)
	Gastrointestinal	4484 (34%)	74 (1.8%)
	Hematology and	224 (1.7%)	61 (1.5%)
	Immunodeficiency		
	Metabolic	1125 (8.5%)	224 (5.6%)
	Congenital or	3642 (27.6%)	896 (22.3%)
	genetic defect		
	Malignancy	158 (1.2%)	68 (1.7%)
	Technology	6432 (48.8%)	849 (21.2%)
	Dependence		
	Transplant	213 (1.6%)	106 (2.6%)

		Prior to Exclusions	Post Exclusions
	Any	12435 (94.3%)	3734 (93%)
Length of Stay:		3 [1, 7]	2 [1, 6]
Median [IQR]			
Length of Stay	a. 0–1 days	4007 (30.4%)	1579 (39.3%)
	b. 2–3 days	3071 (23.3%)	811 (20.2%)
	c. 4–6 days	2367 (17.9%)	697 (17.4%)
	d. 7 days	3744 (28.4%)	927 (23.1%)
Decreased left ventricular function		2419 (18.3%)	1050 (26.2%)
Arrhythmia		938 (7.1%)	301 (7.5%)
Respiratory support		4944 (37.5%)	853 (21.3%)

Table 4A:

Bivariate Analysis for Decreased Left Ventricular (LV) Function

	Decreased LV Function		p
	No	Yes	
N Discharges	2964 (73.8%)	1050 (26.2%)	
Length of Stay (days)	2 [1, 5] ^a	5 [2,10]	<.001
Cost (dollars)	6119 [3317, 17275]	16162 [7038, 45282]	<.001
14 Day Readmission	158 (5.3%)	61 (5.8%)	0.557
Mortality	29 (1%)	61 (5.8%)	<0.001

^a Interquartile range

Table 4B:**Bivariate Analysis for Arrhythmia**

	Arrhythmia		p
	No	Yes	
N Discharges	3713 (92.5%)	301 (7.5%)	
Length of Stay (days)	2 [1, 6]	5 [2, 13]	<.001
Cost (dollars)	7324 [3585, 20906]	25189 [9330, 61637]	<.001
14 Day Readmission	195 (5.3%)	24 (8%)	0.046
Mortality	45 (1.2%)	45 (15%)	<0.001

Table 4C:

Bivariate analysis for respiratory disease

	Respiratory Disease		p
	No	Yes	
N Discharges	3161 (78.7%)	853 (21.3%)	
Length of Stay (days)	2 [1, 5]	6 [3, 13]	<.001
Cost (dollars)	6161 [3390, 15860]	22222 [9024, 61141]	<.001
14 Day Readmission	178 (5.6%)	41 (4.8%)	0.347
Mortality	30 (0.9%)	60 (7%)	<0.001

Table 4D:

Number of Discharges with Each Combination of Exposure

Decreased left ventricular function/cardiomyopathy	Respiratory		N	%
	Arrhythmia	support		
No	No	No	2440	60.8
No	No	Yes	416	10.3
No	Yes	No	72	1.8
No	Yes	Yes	36	0.9
Yes	No	No	541	13.5
Yes	No	Yes	316	7.9
Yes	Yes	No	108	2.7
Yes	Yes	Yes	85	2.1

Table 5A:

Generalized linear mixed effects model with LOS as outcome measure

		LOS	
		Rate Ratio (95% CI)	p
Age	1–4	Reference	
	5–9	1 (0.8,1.1)	0.629
	10–14	1.2 (1,1.4)	0.034
	15–18	1.4 (1.1,1.6)	0.001
	19+	1.4 (1.2,1.7)	<.001
Decreased LV function	Yes	1.4 (1.3,1.5)	<.001
	No	Reference	
Arrhythmia	Yes	1.2 (1.1,1.4)	0.004
	No	Reference	
Respiratory support	Yes	1.6 (1.5,1.8)	<.001
	No	Reference	

Table 5B:

Generalized linear mixed effects model with cost of hospitalization as outcome measure

		Cost	
		Rate Ratio (95% CI)	p
Age	1–4	Reference	
	5–9	1.1 (0.9,1.3)	0.324
	10–14	1.5 (1.3,1.8)	<.001
	15–18	1.7 (1.5,2)	<.001
	19+	1.9 (1.6,2.3)	<.001
Decreased LV function	Yes	1.2 (1.1,1.3)	<.001
	No	Reference	
Arrhythmia	Yes	1.4 (1.2,1.6)	<.001
	No	Reference	
Respiratory support	Yes	1.4 (1.3,1.5)	<.001
	No	Reference	

Table 5C:

Generalized estimating equation with 14-day readmission as outcome measure

		14-Day Readmission	
		Odds Ratio (95% CI)	p
Age	1–4	Reference	
	5–9	0.8 (0.3, 1.9)	0.598
	10–14	0.7 (0.3, 1.3)	0.207
	15–18	0.8 (0.4, 1.6)	0.567
	19+	1.4 (0.5, 3.6)	0.494
Decreased LV function	Yes	1 (0.7, 1.3)	0.780
	No	Reference	
Arrhythmia	Yes	1.5 (0.9, 2.5)	0.128
	No	Reference	
Respiratory support	Yes	0.6 (0.3, 1)	0.045
	No	Reference	